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SEARCH REQUEST FORM
(STIC)

Requestor's Name: David Lukton Examiner number: 71263 Date: 10/6/06

Art Unit: 1654 Phone number: 571-272-0952 Serial Number: 09-854816

Mail Box: 3-C-18 Examiner Rm: 3-D-19 Results format: paper

* * * * * *

Title: CONSTRAINED HELICAL PEPTIDES AND METHODS OF MAKING SAME

Applicants: BRAISTED, ANDREW C.; JUDICE, J. KEVIN; MCDOWELL, ROBERT S.; PHELAN, J. CHRISTOPHER; STAROVASNIK, MELISSA A.; WELLS, JAMES A.;

Earliest Priority Date: 11/6/96

* * * *

Applicants are claiming cyclic peptides as shown on the attached sheet.

R^1 = anything;

R^2 = anything;

R^3 = anything;

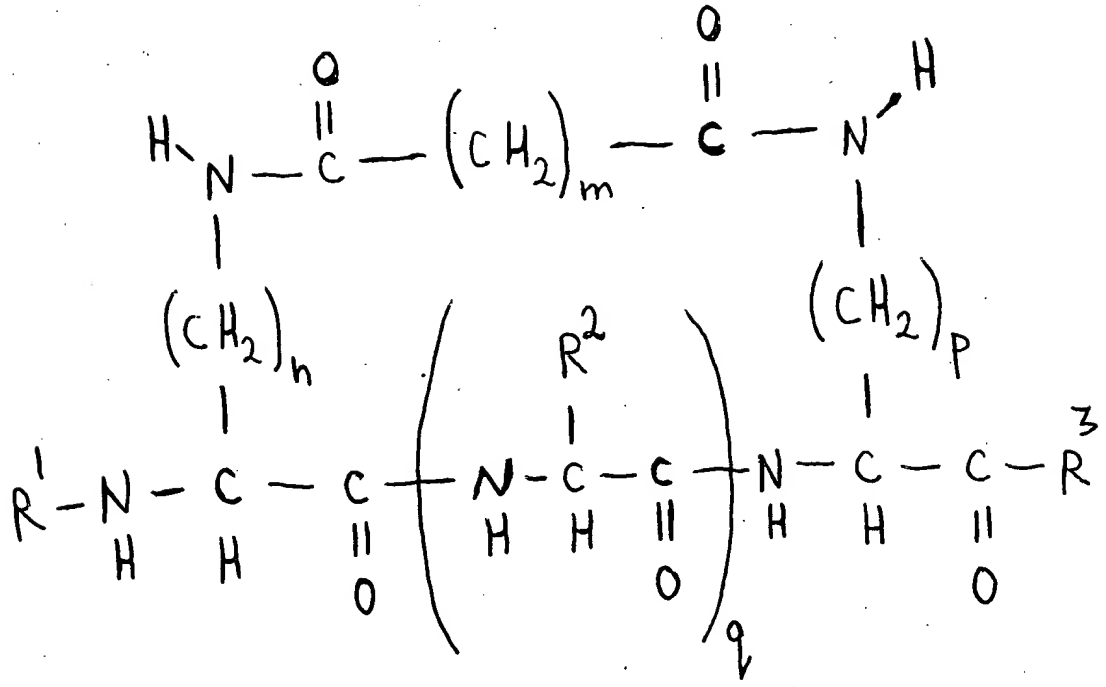
n = an integer of 3 - 4

p = an integer of 3 - 4

m = an integer of 1 - 6

q = an integer of 6 (no more and no less)

09/854816



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DICTIONARY FILE UPDATES:    9 OCT 2006  HIGHEST RN 910025-51-3

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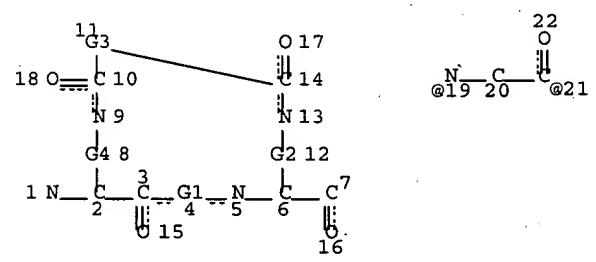


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FILE COVERS 1907 - 10 Oct 2006 VOL 145 ISS 16
FILE LAST UPDATED: 8 Oct 2006 (20061008/ED)

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This file contains CAS Registry Numbers for easy and accurate substance identification.

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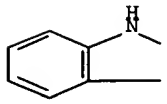
L14 ANSWER 1 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN
AN 2002:251292 HCAPLUS Full-text
DN 137:211058
TI Exploration of the DTrp-NMeLys Motif in the Search for potent somatostatin antagonists
AU Rajeswaran, W. G.; Murphy, William A.; Taylor, John E.; Coy, David H.
CS Department of Medicine, SL 53, Peptide Research Labs, Tulane University Health Sciences Center, New Orleans, LA, 70112, USA
SO Bioorganic & Medicinal Chemistry (2002), 10(6), 2023-2029
CODEN: BMECEP; ISSN: 0968-0896
PB Elsevier Science Ltd.
DT Journal
LA English
AB Previous studies from this laboratory demonstrated that N-methylation at Lys5 residue in somatostatin octapeptide antagonist analogs increased the GH release inhibition potency by as much as 300%. The authors have now further investigated N-methylation of this Lys5 residue in conjunction with a number of N- and C-terminal modifications previously found to give highly potent somatostatin receptor antagonists. Synthetic analogs were tested in a functional assay for their ability to inhibit somatostatin-inhibited GH release from rat pituitary cells in culture and to displace 125I-labeled somatostatin from CHO cells transfected with the five known human somatostatin receptors. Several interesting observations resulted from the study. Replacement of lipophilic Nal8 at the C-terminus with a hydrophilic His8 resulted in the increased affinity and selectivity for type 2 receptor to give the most potent antagonist analog yet discovered (Ki, 1.5 nM), although in the rat pituitary cells inhibitory activity on somatostatin inhibited GH release decreased somewhat. A His3 substitution within the cyclic portion of the analogs retained pituitary cell potency and affinity for type 2 receptor as did substitution with Bip8 and Fpa1. Replacement of Cpa1 with Iph1 did not effect the affinity for type 2 receptor significantly, but did decrease the effects on rat cell GH release. Iph3 within-ring substitution increased the selectivity for sst2 appreciably although the affinity for that receptor was considerably decreased. Substitution of Npa3 resulted in good selectivity for sst2 receptor. Replacement of Nal8 with D-Trp8 also increased the selectivity for type 2 receptor. Use of a 'bivalent ligand' approach in which two peptides were joined by 4,4'-biphenyldicarbonyl as a spacer destroyed the affinity for all the subtypes, however, the bivalent ligand formed with the Ahp spacer displayed significant affinity and high selectivity for the type 2 receptor.
IT **455333-34-3P**
RL: PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)
(exploration of DTrp-NMeLys motif in search for potent human somatostatin receptor antagonists)
RN 455333-34-3 HCAPLUS
CN L-Alaninamide, 3-[(3-carboxy-1-oxopropyl)amino]-L-alanyl-D-cysteinyl-L-tyrosyl-D-tryptophyl-N2-methyl-L-lysyl-L-threonyl-L-cysteinyl-3-amino-,

(1→8)-lactam (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



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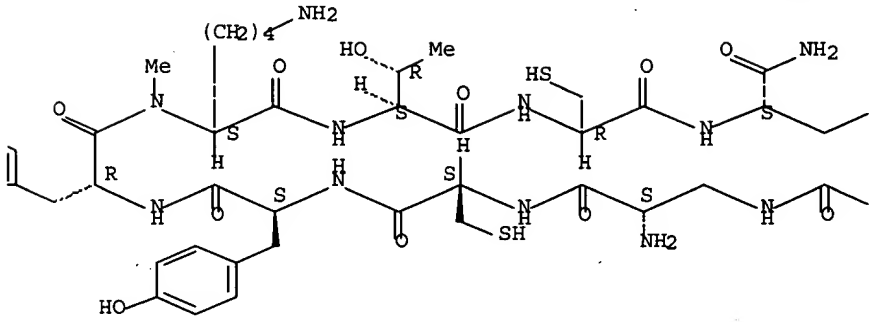

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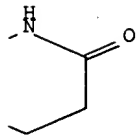

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Yamada, Y	1992	89	251	Proc Natl Acad Sci U	HCAPLUS
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L14 ANSWER 2 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN
AN 2001:573536 HCAPLUS Full-text
DN 135:122757
TI Preparation of constrained helical peptides
IN Braisted, Andrew C.; Judice, J. Kevin; Mcdowell, Robert S.; Phelan, J. Christopher; Starovasnik, Melissa A.; Wells, James A.
PA Genentech, Inc., USA
SO U.S., 175 pp., Cont.-in-part of U.S. Ser. No. 876,698, abandoned.
CODEN: USXXAM
DT Patent
LA English
FAN.CNT 149

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AB Cyclic peptides, e.g., [NHCO(CH2)mCH(NHX)CO-Z-NHCH(COYS)(CH2)pCONH](CH2)n [(CH2)n is attached to NH end groups, S is absent or is a macromol., X is H or is any amino acid or amino acid sequence, Y is absent or is hydroxyl if S is absent or is any amino acid or amino acid sequence, Z is any amino acid sequence consisting of six amino acids, m and p are 0-6, n is an integer greater than zero], with constrained region(s) having an α-helical conformation, were prepared Constrained helical peptides having amino acid sequences from HIV gp41 are provided, as is their use in preparing antibodies that prevent viral membrane fusion. Thus, cyclic peptide FNM(5)QQRRFY(6)ALH (5 and 6 represent glutamic acid residues cyclized via 1,5-pentanediamine) was prepared by standard solid phase protocols.

IT **137363-78-1P 185335-95-9P**

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation)

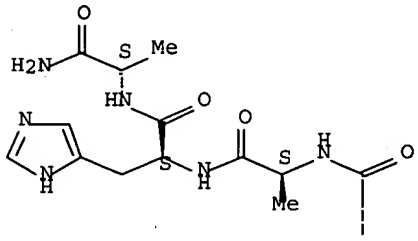
(preparation of constrained helical peptides)

RN 137363-78-1 HCAPLUS

CN L-Alaninamide, N-acetyl-L-alanyl-L-α-glutamyl-6-mercapto-D-norleucyl-L-alanyl-L-alanyl-L-alanyl-L-lysyl-L-phenylalanyl-L-leucyl-6-mercapto-L-norleucyl-L-alanyl-L-histidyl-, cyclic (3→10)-disulfide (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



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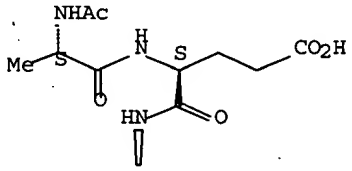

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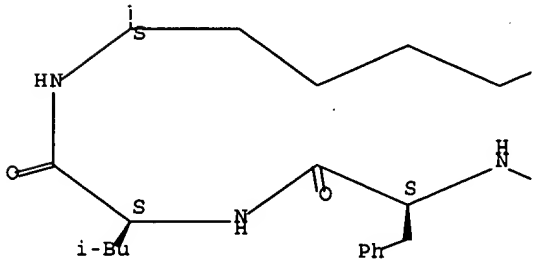

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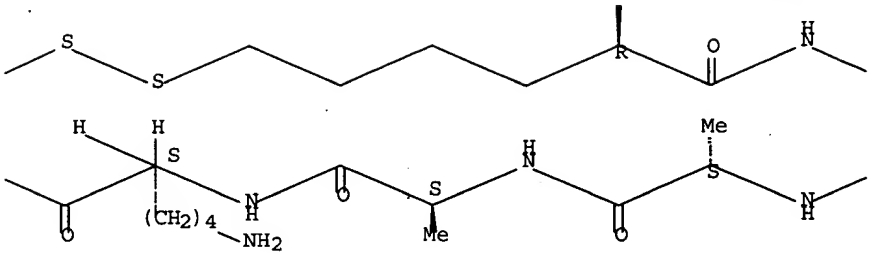

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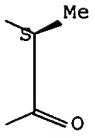
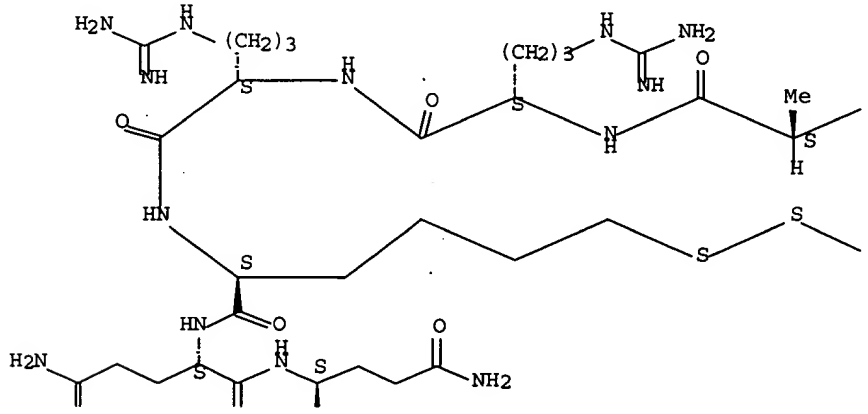


RN 185335-95-9 HCAPLUS

CN L-Glutamamide, N-acetyl-L-threonyl-L-asparaginyl-6-mercapto-D-norleucyl-L-α-aspartyl-L-leucyl-L-alanyl-L-alanyl-L-arginyl-L-arginyl-6-mercapto-L-norleucyl-L-glutaminyl-, cyclic (3→10)-disulfide (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



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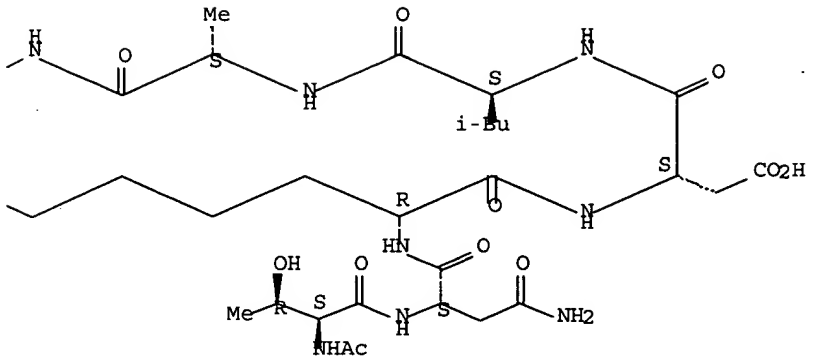

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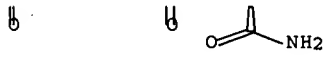

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TI Preparation of constrained helical peptides
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PA Genentech, Inc., USA
SO PCT Int. Appl., 281 pp.
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DT Patent
LA English
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2002US-0211858	A1	20020802
2003AU-0261484	A	20031106
2004US-0797366	A1	20040309

AB Cyclic peptides, e.g., [NHCO(CH2)mCH(NHX)CO-Z-NHCH(COYS)(CH2)pCONH](CH2)n [(CH2)n is attached to NH end groups, S is absent or is a macromol., X is H or is any amino acid or amino acid sequence, Y is absent or is hydroxyl if S is absent or is any amino acid or amino acid sequence, Z is any amino acid sequence consisting of six amino acids, m and p are 0-6, n is an integer greater than zero], with constrained region(s) having an α-helical conformation, were prepared Constrained helical peptides having amino acid sequences from HIV gp41 are provided, as is their use in preparing antibodies that prevent viral membrane fusion. Thus, cyclic peptide FNM(5)QQRRFY(6)ALH (5 and 6 represent glutamic acid residues cyclized via 1,5-pentanediamine) was prepared by standard solid phase protocols.

IT **137363-78-1P 185335-95-9P**
RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation)
(preparation of constrained helical peptides)

RN 137363-78-1 HCAPLUS

CN L-Alaninamide, N-acetyl-L-alanyl-L-α-glutamyl-6-mercapto-D-norleucyl-L-alanyl-L-alanyl-L-alanyl-L-lysyl-L-phenylalanyl-L-leucyl-6-mercapto-L-norleucyl-L-alanyl-L-histidyl-, cyclic (3→10)-disulfide (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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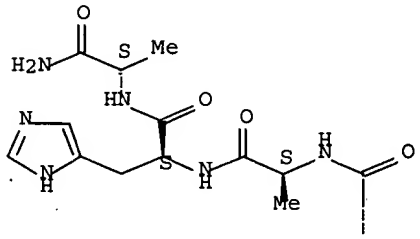

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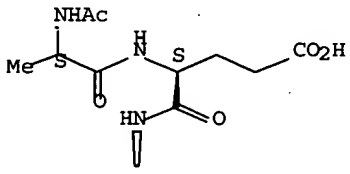

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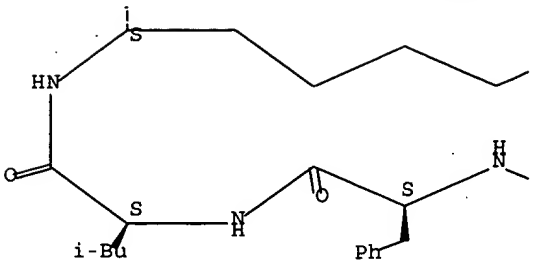

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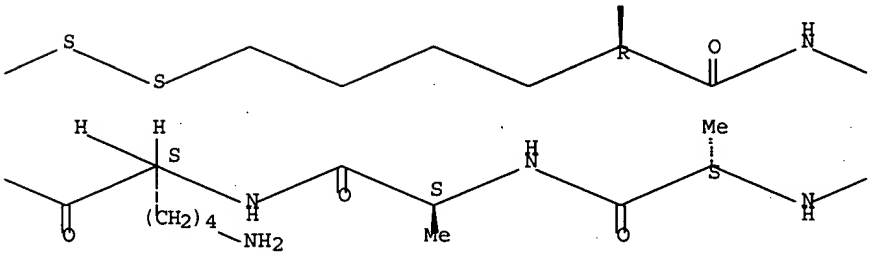


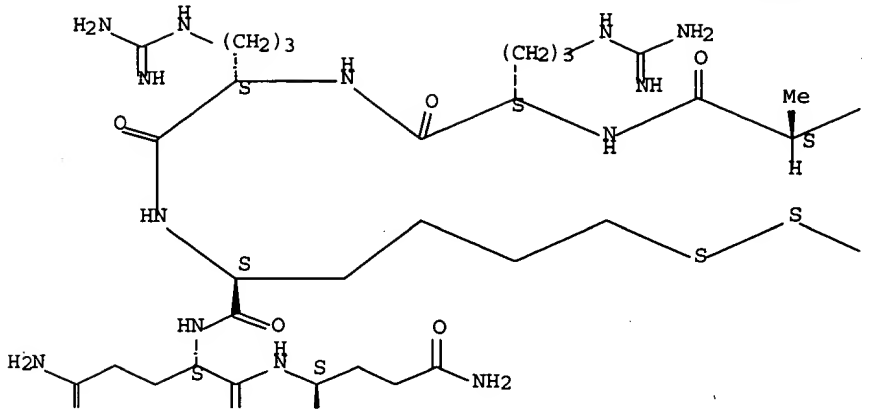
PAGE 2-C

RN 185335-95-9 HCAPLUS

CN L-Glutamamide, N-acetyl-L-threonyl-L-asparaginyl-6-mercapto-D-norleucyl-L-α-aspartyl-L-leucyl-L-alanyl-L-alanyl-L-arginyl-L-arginyl-6-mercapto-L-norleucyl-L-glutaminyl-, cyclic (3→10)-disulfide (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



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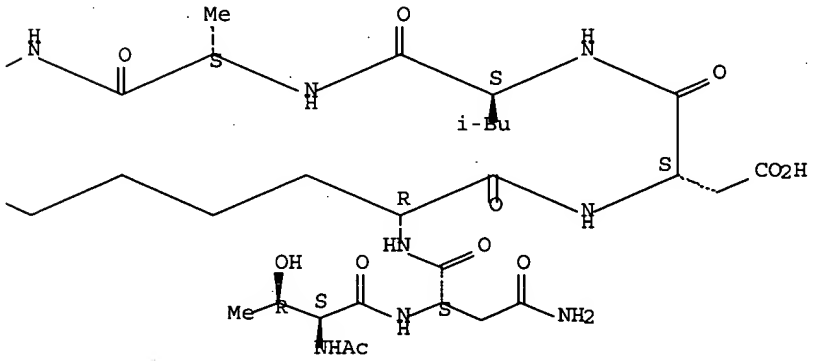

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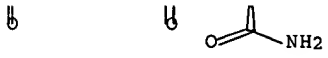

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Referenced Author (RAU)	Year (RPY)	VOL (RVL)	PG (RPG)	Referenced Work (RWK)	Referenced File
Genentech Inc	1991			WO---9115512 A	HCAPLUS
Lawless, M	1996	35	13697	BIOCHEMISTRY	HCAPLUS
Phelan, J	1997	119	455	JOURNAL OF THE AMERI	HCAPLUS
Smithkline Beecham Corp	1992			WO---9209625 A	HCAPLUS
Univ Pennsylvania	1995			WO---9534312 A	HCAPLUS
Zhang, X	1997	15	150	NATURE BIOTECHNOLOGY	HCAPLUS

L14 ANSWER 4 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN
AN 1997:80130 HCAPLUS Full-text
DN 126:75230
TI A General Method for Constraining Short Peptides to an α-Helical Conformation
AU Phelan, J. Christopher; Skelton, Nicholas J.; Braisted, Andrew C.;

McDowell, Robert S.
CS Department of Bioorganic Chemistry and Department of Protein Engineering, Genentech Inc., South San Francisco, CA, 94080, USA
SO Journal of the American Chemical Society (1997), 119(3), 455-460
CODEN: JACSAT; ISSN: 0002-7863
PB American Chemical Society
DT Journal
LA English
AB A method for constraining short peptides (containing fewer than 20 residues) of an arbitrary sequence to an α-helical conformation (.apprx.100% helical in H2O at 25 °C) is presented. Gln residues at positions i and i + 7 of the peptides were tethered with an alkanediyl chain between the side chain nitrogen atoms. Peptides containing this tether were readily synthesized on the solid phase by amide formation between an α,ω-diaminoalkane and the side chain carboxylates of Glu residues. The resulting cyclic peptides were studied by NMR and CD and were found to adopt an α-helical conformation in aqueous solution and this α-helix was thermally stable to ≥40°. Corresponding untethered control peptides with N-methylglutamine at the i and i + 7 positions lacked helicity under the same conditions.
IT **137363-78-1P 185335-95-9P**
RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation)
(preparation of short peptides constrained to an α-helical conformation)
RN 137363-78-1 HCAPLUS
CN L-Alaninamide, N-acetyl-L-alanyl-L-α-glutamyl-6-mercapto-D-norleucyl-L-alanyl-L-alanyl-L-alanyl-L-lysyl-L-phenylalanyl-L-leucyl-6-mercapto-L-norleucyl-L-alanyl-L-histidyl-, cyclic (3→10)-disulfide (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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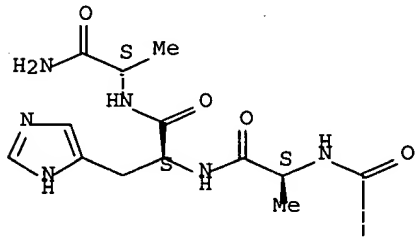

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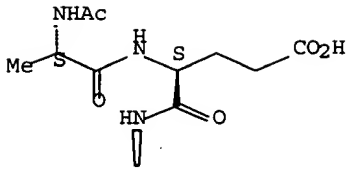

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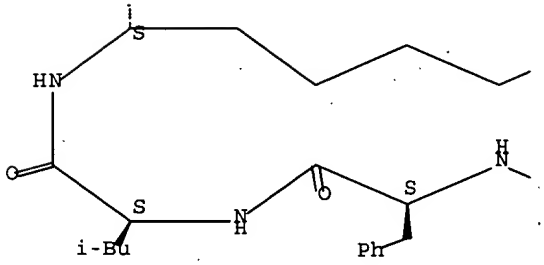

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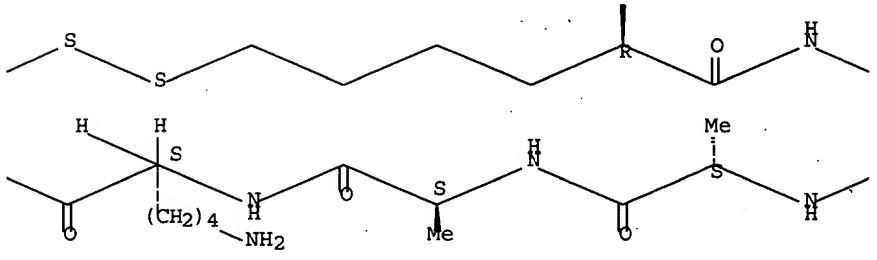

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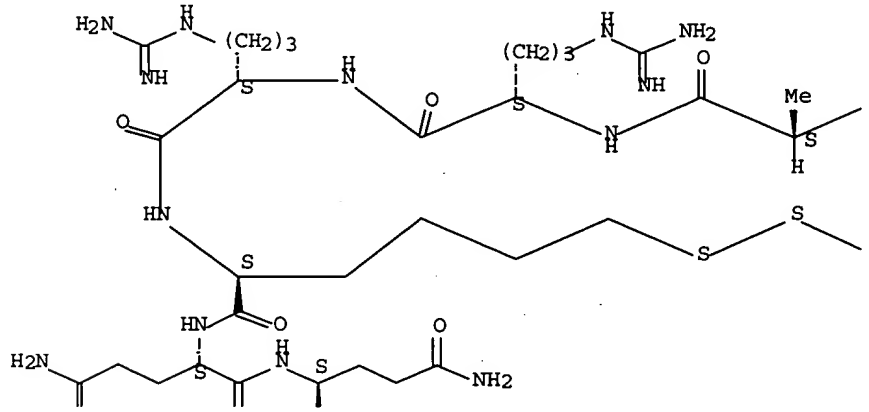
RN 185335-95-9 HCAPLUS
CN L-Glutamamide, N-acetyl-L-threonyl-L-asparaginyl-6-mercapto-D-norleucyl-L-

α-aspartyl-L-leucyl-L-alanyl-L-alanyl-L-arginyl-L-arginyl-6-mercapto-L-norleucyl-L-glutaminyl-, cyclic (3→10)-disulfide (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



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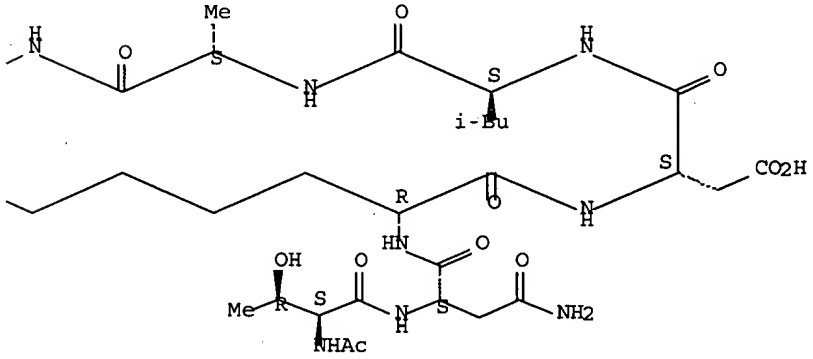

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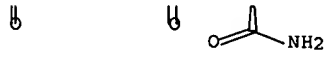

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Bernstein, F	1977	112	535	J Mol Biol	HCAPLUS
Bierzynski, A	1982	79	2470	Proc Natl Acad Sci U	HCAPLUS
Blaney, J				DGEOM, QCPE No 590	
Bracken, C	1994	116	6431	J Am Chem Soc	HCAPLUS
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Brown, J	1971	10	470	Biochemistry	HCAPLUS

Callewaert, G	1968	1	111	FEBS Lett	HCAPLUS
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Fairman, R	1992	114	5458	J Am Chem Soc	HCAPLUS
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Habermann, E	1965	343	192	Biochem Z	HCAPLUS
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Todd, R	1991	10	156	Proteins:Struct, Fun	HCAPLUS
Wuthrich, K	1986			NMR of Proteins and	
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L14 ANSWER 5 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN
AN 1991:680537 HCAPLUS Full-text
DN 115:280537
TI General approach to the synthesis of short α-helical peptides
AU Jackson, David Y.; King, David S.; Chmielewski, Jean; Singh, Sunil; Schultz, Peter G.
CS Dep. Chem., Univ. California, Berkeley, CA, 94720, USA
SO Journal of the American Chemical Society (1991), 113(24), 9391-2
CODEN: JACSAT; ISSN: 0002-7863
DT Journal
LA English
OS CASREACT 115:280537
AB Short peptides have been synthesized which contain a single intramol. disulfide bond that stabilizes two helical turns. Peptides containing D- and L-S-(acetamidomethyl)-2-amino-6-mercaptohexanoic acid at the i and i + 7 residue, resp., show only slight α-helicity in the reduced form in aqueous solution On oxidation, these peptides exhibit a large increase in α-helicity in water both at 0° and 60°. This approach has been used to generate eight and sixteen amino acid peptides with high helicity. Oxidation can be carried out under a wide variety of conditions with peptides that contain a large variety of functional groups.
IT **137363-78-1P**
RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation)
(preparation and conformation of)
RN 137363-78-1 HCAPLUS
CN L-Alaninamide, N-acetyl-L-alanyl-L-α-glutamyl-6-mercapto-D-norleucyl-L-alanyl-L-alanyl-L-alanyl-L-lysyl-L-phenylalanyl-L-leucyl-6-mercapto-L-

norleucyl-L-alanyl-L-histidyl-, cyclic (3→10)-disulfide (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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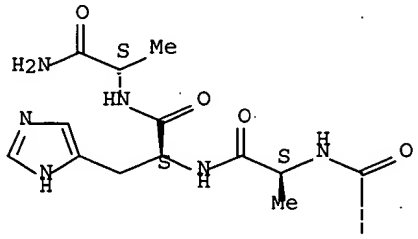

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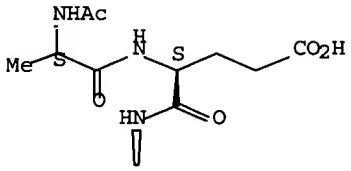

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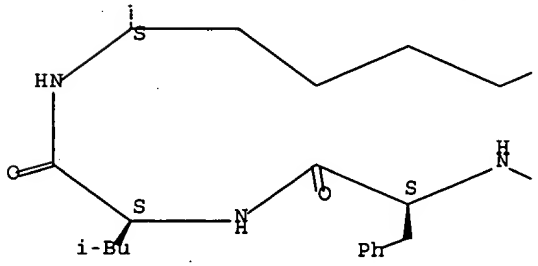

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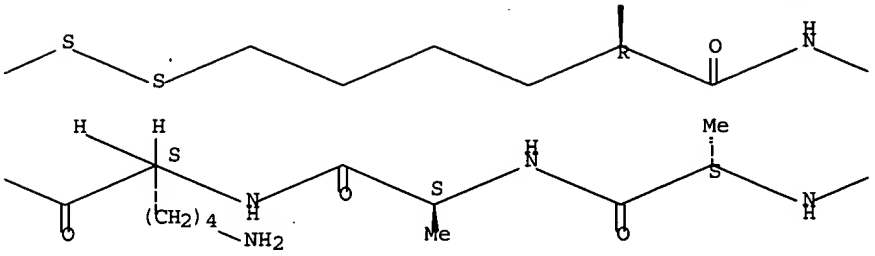

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FILE 'USPAT2' ENTERED AT 09:44:57 ON 10 OCT 2006
CA INDEXING COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

=> d bib abs fhitstr hitrn l16 tot

L16  ANSWER 1 OF 2  USPATFULL on STN
AN      2003:244455  USPATFULL  Full-text
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        Desnoyers, Luc, San Francisco, CA, UNITED STATES
        Ferrara, Napoleone, San Francisco, CA, UNITED STATES
        Fong, Sherman, Alameda, CA, UNITED STATES
        Gao, Wei-Qiang, Palo Alto, CA, UNITED STATES
        Goddard, Audrey, San Francisco, CA, UNITED STATES
        Gurney, Austin L., Belmont, CA, UNITED STATES
        Pan, James, Belmont, CA, UNITED STATES
        Roy, Margaret Ann, San Francisco, CA, UNITED STATES
        Stewart, Timothy A., San Francisco, CA, UNITED STATES
        Tumas, Daniel, Orinda, CA, UNITED STATES
        Watanabe, Colin K., Moraga, CA, UNITED STATES
        Wood, William I., Hillsborough, CA, UNITED STATES
PA      GENENTECH, INC. (U.S. corporation)
PI      US2003170864       A1   20030911
AI      2001US-0866034     A1   20010525 (9)
RLI     Continuation of Ser. No. 2000WO-US14941, filed on 30 May 2000, UNKNOWN
        Continuation of Ser. No. 2000WO-US15264, filed on 2 Jun 2000, UNKNOWN
        Continuation of Ser. No. 2000WO-US32678, filed on 1 Dec 2000, UNKNOWN
DT      Utility
FS      APPLICATION
LREP    KNOBBE, MARTENS, OLSON AND BEAR, LLP, 2040 MAIN STREET, FOURTEENTH
        FLOOR, IRVINE, CA, 92614
CLMN    Number of Claims: 21
ECL     Exemplary Claim: 1
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DRWN 18 Drawing Page(s)
LN.CNT 7716
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention is directed to novel polypeptides and to nucleic acid molecules encoding those polypeptides. Also provided herein are vectors and host cells comprising those nucleic acid sequences, chimeric polypeptide molecules comprising the polypeptides of the present invention fused to heterologous polypeptide sequences, antibodies which bind to the polypeptides of the present invention and to methods for producing the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

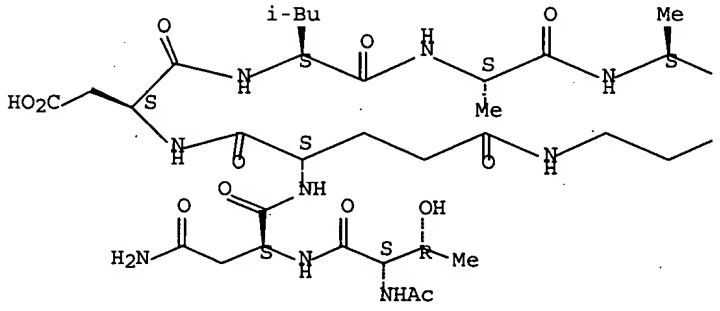
IT **185335-87-9P**
(preparation of constrained helical peptides)

RN 185335-87-9 USPATFULL

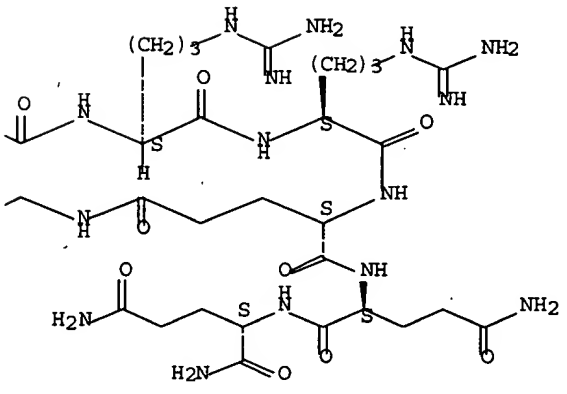
CN L-Glutamamide, N-acetyl-L-threonyl-L-asparaginyl-N-(3-aminopropyl)-L-glutaminyl-L-α-aspartyl-L-leucyl-L-alanyl-L-alanyl-L-arginyl-L-arginyl-L-α-glutamyl-L-glutaminyl-, (10→3)-lactam (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



PAGE 1-B



IT **185335-87-9P 185335-88-0P 185335-89-1P 185335-91-5P 185335-92-6P 185335-93-7P 207676-47-9P**
(preparation of constrained helical peptides)

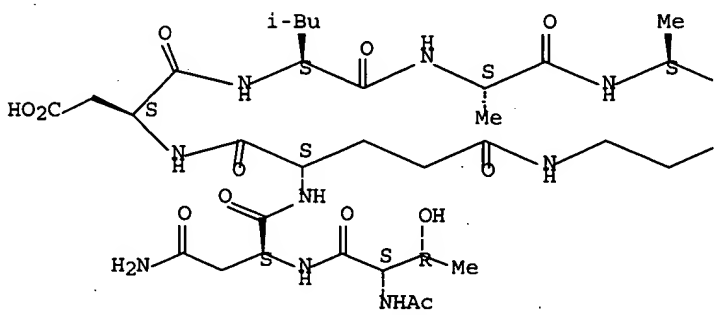
L16 ANSWER 2 OF 2 USPATFULL on STN
AN 2001:125963 USPATFULL Full-text
TI Constrained helical peptides and methods of making same
IN Braisted, Andrew C., San Francisco, CA, United States
Judice, J. Kevin, San Francisco, CA, United States
McDowell, Robert S., San Francisco, CA, United States
Phelan, J. Christopher, San Francisco, CA, United States
Starovasnik, Melissa A., Burlingame, CA, United States
Wells, James A., Burlingame, CA, United States
PA Genentech, Inc., South San Francisco, CA, United States (U.S. corporation)
PI US---6271198 B1 20010807
AI 1997US-0965056 19971105 (8)
RLI Continuation-in-part of Ser. No. 1997US-0876698, filed on 16 Jun 1997, now abandoned , said Ser. No. US 965056 And Ser. No. 1996US-0743698, filed on 6 Nov 1996
PRAI 1997US-049787P 19970616 (60)
DT Utility
FS GRANTED
EXNAM Primary Examiner: Jones, Dwayne C.; Assistant Examiner: Delacroix-Muirheid, C.
LREP Piper Marbury Rudnick & Wolfe LLP, Kelber, Steven B.
CLMN Number of Claims: 4
ECL Exemplary Claim: 1
DRWN 40 Drawing Figure(s); 34 Drawing Page(s)
LN.CNT 6260
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB Provided are cyclized peptides with a constrained region(s) having an α-helical conformation. Constrained helical peptides having amino acid sequences from HIV gp41 are provided, as is their use in preparing antibodies that prevent viral membrane fusion. Also provided are methods for making such cyclized peptides.

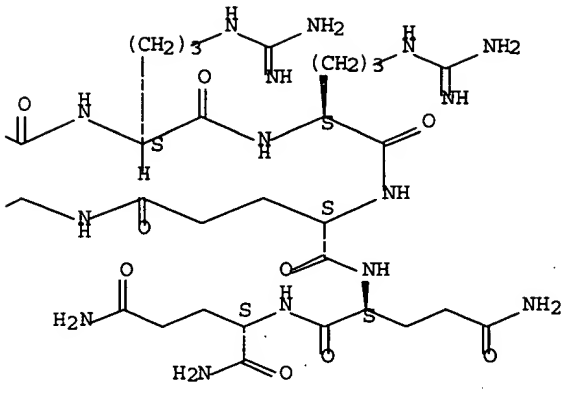
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
IT **185335-87-9P**
(preparation of constrained helical peptides)
RN 185335-87-9 USPATFULL
CN L-Glutamamide, N-acetyl-L-threonyl-L-asparaginyl-N-(3-aminopropyl)-L-glutaminyl-L-α-aspartyl-L-leucyl-L-alanyl-L-alanyl-L-arginyl-L-arginyl-L-α-glutamyl-L-glutaminyl-, (10→3)-lactam (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



PAGE 1-B



IT 185335-87-9P 185335-88-0P 185335-89-1P
185335-91-5P 185335-92-6P 185335-93-7P
207676-47-9P
(preparation of constrained helical peptides)

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